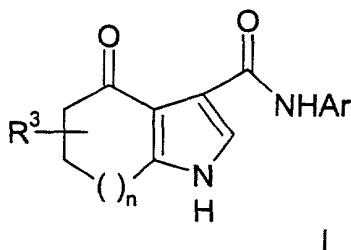
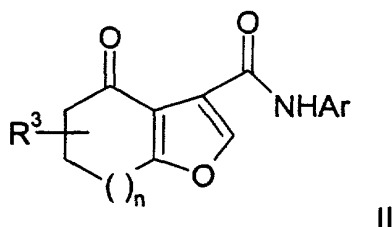


We claim:

1. A method of preparing a compound of the formula:



comprising reacting a compound of the formula:



with an excess of ammonia source in a reaction inert solvent at an elevated temperature until reaction is complete;

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH<sub>2</sub>)<sub>m</sub>-NR<sup>1</sup>R<sup>2</sup>, -O(CH<sub>2</sub>)C(O)OR<sup>4</sup>, -CH(NR<sup>7</sup>R<sup>8</sup>)CH<sub>3</sub>, -CH<sub>2</sub>CH(NR<sup>5</sup>R<sup>6</sup>)CH<sub>3</sub>, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C<sub>1</sub> - C<sub>6</sub> alkoxy, C<sub>1</sub> - C<sub>6</sub> alkyl, C<sub>2</sub> - C<sub>6</sub> alkenyl, C<sub>1</sub> - C<sub>6</sub> perfluoroalkyl, F, Cl, and Br, wherein:

R<sup>1</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup> and R<sup>7</sup> are independently selected from hydrogen and C<sub>1</sub> - C<sub>6</sub> alkyl;

R<sup>2</sup>, R<sup>6</sup>, and R<sup>8</sup> are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

2. The method of claim 1 wherein Ar is phenyl substituted with said one or two groups.

3. The method of claim 1 wherein said nitrogen protecting group is -C(O)C<sub>1</sub>-C<sub>6</sub> alkoxy.

4. The method of claim 1 wherein said nitrogen protecting group is benzyloxycarbonyl, fluorenyloxycarbonyl, acetyl, trifluoroacetyl, chloroacetyl, benzoyl, t-butyloxycarbonyl, or benzyl.

5. The method of claim 1 wherein said compound of formula I is selected from the group consisting of

Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;



[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

Butyl-(2-[5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy]-ethyl)-carbamic acid tert-butyl ester;

5        4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

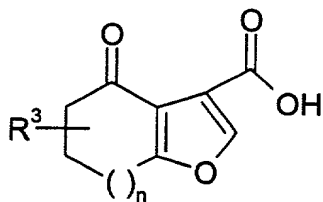
(1-Methyl-2-[4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl]-ethyl)-carbamic acid tert-butyl ester;

10        (2-[4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy]-ethyl)-propyl-carbamic acid tert-butyl ester; and

{2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

6.        A method according to claim 1 further wherein said compound of formula II is prepared by

15        (a)        reacting a compound of the formula



III

with an excess of an acid chloride or anhydride in a reaction inert solvent containing an excess of an acid acceptor until reaction is complete; and

20        (b)        adding an equivalent amount of  $\text{NH}_2\text{-Ar}$  to the solution of step (a) and holding until reaction is complete.

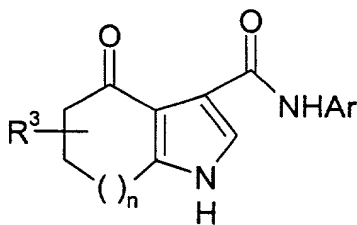
7.        The method of claim 6 wherein said acid chloride is ethylchloroformate.

8.        The method according to claim 1 which further comprises removing said nitrogen protecting group.

25        9.        The method according to claim 5 which further comprises removing said nitrogen protecting group.

10.        A compound of the following formula:





I

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH<sub>2</sub>)<sub>m</sub>-NR<sup>1</sup>R<sup>2</sup>, -O(CH<sub>2</sub>)<sub>l</sub>C(O)OR<sup>4</sup>, -CH(NR<sup>7</sup>R<sup>8</sup>)CH<sub>3</sub>, -CH<sub>2</sub>CH(NR<sup>5</sup>R<sup>6</sup>)CH<sub>3</sub>, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C<sub>1</sub> - C<sub>6</sub> alkoxy, C<sub>1</sub> - C<sub>6</sub> alkyl, C<sub>2</sub> - C<sub>6</sub> alkenyl, C<sub>1</sub> - C<sub>6</sub> perfluoroalkyl, F, Cl, and Br, wherein:

R<sup>1</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup> and R<sup>7</sup> are independently selected from hydrogen and C<sub>1</sub> - C<sub>6</sub> alkyl;

R<sup>2</sup>, R<sup>6</sup>, and R<sup>8</sup> are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

11. A compound of claim 10 selected from the group consisting of:

Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy}-ethyl)-carbamic acid tert-butyl ester;

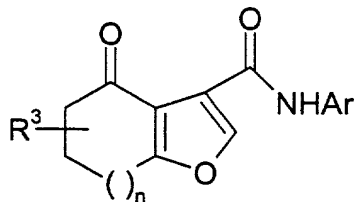
4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

(1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

(2-{4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and

{2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

12. A compound of the following formula:



II



wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with  $-O-(CH_2)_m-NR^1R^2$ ,  $-O(CH_2)C(O)OR^4$ ,  $-CH(NR^7R^8)CH_3$ ,  $-CH_2CH(NR^5R^6)CH_3$ , or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from  $C_1 - C_6$  alkoxy,  $C_1 - C_6$  alkyl,  $C_2 - C_6$  alkenyl,  $C_1 - C_6$  perfluoroalkyl, F, Cl, and Br, wherein:

- 5         $R^1$ ,  $R^3$ ,  $R^4$ ,  $R^5$  and  $R^7$  are independently selected from hydrogen and  $C_1 - C_6$  alkyl;  
          $R^2$ ,  $R^6$ , and  $R^8$  are independently selected from nitrogen protecting groups;  
         m and l are integers independently selected from 1 to 6; and  
         n is an integer from 0 to 2.

13.        The compound of claim 12 selected from the group consisting of:

- 10        Methyl-(1-{4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenyl}-ethyl)-  
         carbamic acid tert-butyl ester;  
         [2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenoxy)-  
         ethyl]-propyl-carbamic acid tert-butyl ester;  
         Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-pyridin-2-yloxy}-  
15        ethyl)-carbamic acid tert-butyl ester;  
         4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]furan-3-carboxylic acid (2-fluoro-4-hydroxy-  
         phenyl)-amide;  
         (1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenyl}-  
         ethyl)-carbamic acid tert-butyl ester;  
20        (2-{4-[(4-Oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy}-ethyl)-  
         propyl-carbamic acid tert-butyl ester; and  
         {2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy}-acetic  
         acid ethyl ester.